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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/583,640	06/20/2006	Koen De Keersmaecker	IMEC320.001APC	3620
20995 7590 05/04/2011 KNOBBE MARTENS OLSON & BEAR LLP 2040 MAIN STREET			EXAMINER	
			SAKELARIS, SALLY A	
FOURTEENTH FLOOR IRVINE, CA 92614			ART UNIT	PAPER NUMBER
			1773	
			NOTIFICATION DATE	DELIVERY MODE
			05/04/2011	ELECTRONIC

# Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

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	Application No.	Applicant(s)	
	10/583,640	KEERSMAECKER ET AL.	
Office Action Summary	Examiner	Art Unit	
	SALLY A. SAKELARIS	1773	
The MAILING DATE of this communication a Period for Reply	appears on the cover sheet wit	h the correspondence address	
A SHORTENED STATUTORY PERIOD FOR REF WHICHEVER IS LONGER, FROM THE MAILING  - Extensions of time may be available under the provisions of 37 CFR after SIX (6) MONTHS from the mailing date of this communication.  - If NO period for reply is specified above, the maximum statutory period.  - Failure to reply within the set or extended period for reply will, by stat Any reply received by the Office later than three months after the mai earned patent term adjustment. See 37 CFR 1.704(b).	DATE OF THIS COMMUNIC 1.136(a). In no event, however, may a re od will apply and will expire SIX (6) MONT tute, cause the application to become ABA	ATION. ply be timely filed  HS from the mailing date of this communication. NDONED (35 U.S.C. § 133).	
Status			
1) ■ Responsive to communication(s) filed on <u>28</u> 2a) ■ This action is <b>FINAL</b> . 2b) ■ The sum of the su	his action is non-final. vance except for formal matte	•	
Disposition of Claims			
4) ☐ Claim(s) 21-24,26-32,41 and 50-62 is/are per 4a) Of the above claim(s) is/are withdress.  5) ☐ Claim(s) is/are allowed.  6) ☐ Claim(s) 21-24,26-32,41 and 50-62 is/are regress.  7) ☐ Claim(s) 52-62 is/are objected to.  8) ☐ Claim(s) are subject to restriction and application Papers.	rawn from consideration.  ejected.  d/or election requirement.		
9) The specification is objected to by the Exami 10) The drawing(s) filed on is/are: a) a Applicant may not request that any objection to the Replacement drawing sheet(s) including the correctable.  11) The oath or declaration is objected to by the	ccepted or b) objected to be ne drawing(s) be held in abeyand ection is required if the drawing(s	ee. See 37 CFR 1.85(a). s) is objected to. See 37 CFR 1.121(d).	
Priority under 35 U.S.C. § 119			
12) Acknowledgment is made of a claim for foreign a) All b) Some * c) None of:  1. Certified copies of the priority docume 2. Certified copies of the priority docume 3. Copies of the certified copies of the priority docume application from the International Bure * See the attached detailed Office action for a limit	ents have been received. ents have been received in Apriority documents have been reau (PCT Rule 17.2(a)).	oplication No received in this National Stage	
Attachment(s)  1) Notice of References Cited (PTO-892)  2) Notice of Draftsperson's Patent Drawing Review (PTO-948)		ummary (PTO-413) /Mail Date	
Notice of Draftsperson's Patent Drawing Review (PTO-948)     Information Disclosure Statement(s) (PTO/SB/08)     Paper No(s)/Mail Date	_	formal Patent Application	

### **DETAILED ACTION**

### **Continued Examination Under 37 CFR 1.114**

A request for continued examination under 37 CFR 1.114, including the fee set forth in 37 CFR 1.17(e), was filed in this application after final rejection. Since this application is eligible for continued examination under 37 CFR 1.114, and the fee set forth in 37 CFR 1.17(e) has been timely paid, the finality of the previous Office action has been withdrawn pursuant to 37 CFR 1.114. Applicant's submission filed on 3/28/2011 has been entered.

## **Claim Objections**

Claims 52-62 are objected to because of the following informalities: There are 2 claim 52s. Appropriate correction is required.

### Claim Rejections - 35 USC § 112

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

Claims 51-59 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

Claim 51 recites that the "field effect transistor" is "attached upside down to a solid substrate by a bottom auxiliary layer". It is not clear if the applicant is intending to claim that the included circuit is to be built upside-down (i.e. the positive rail is ground), if the solid substrate is right side up or upside down, or if the FET is right side up or upside down. Furthermore the term

"upside down" and "backside" are relative terms depending on one's vantage point. The applicant should clarify the objective positively recited structure of this device. It is presently unclear.

### Claim Rejections - 35 USC § 103

The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

(a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.

The factual inquiries set forth in Graham v. John Deere Co., 383 U.S. 1, 148 USPQ 459 (1966), that are applied for establishing a background for determining obviousness under 35 U.S.C. 103(a) are summarized as follows:

- 1. Determining the scope and contents of the prior art.
- 2. Ascertaining the differences between the prior art and the claims at issue.
- 3. Resolving the level of ordinary skill in the pertinent art.
- 4. Considering objective evidence present in the application indicating obviousness or nonobviousness.

This application currently names joint inventors. In considering patentability of the claims under 35 U.S.C. 103(a), the examiner presumes that the subject matter of the various claims was commonly owned at the time any inventions covered therein were made absent any evidence to the contrary. Applicant is advised of the obligation under 37 CFR 1.56 to point out the inventor and invention dates of each claim that was not commonly owned at the time a later invention was made in order for the examiner to consider the applicability of 35 U.S.C. 103(c) and potential 35 U.S.C. 102(e), (f) or (g) prior art under 35 U.S.C. 103(a).

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1. Claims 21-32, 41, and 50-59 are rejected under 35 U.S.C. 103(a) as being unpatentable over Yousaf et al.(PNAS, 2001) in view of Hollis et al. (US 5653,939)

With regard to claim 21, Yousaf et al. teach a sensing device for sensing a specific binding between an analyte and a recognition molecule, the sensing device comprising: a patterned, localized, and individually addressable microelectronic sensor comprising an individually addressable activation element and a plurality of self-aligned recognition molecules (i.e., the monolayer is interpreted as being self-aligned, localized, and individually addressable in the absence of any structural requirements for these components see Figure 1) microelectronically addressable sensor surface, the sensor surface comprising a recognition molecule (self-assembled monolayer (SAM) of alkanethiolates) and an activation element (i.e., electrical potential) which is a thermal or electrochemical activation element, wherein the activation element is configured to activate the sensor surface by heating or adjusting an oxidation state of a part of the anchoring layer through an applied voltage or current, wherein the recognition molecule (i.e., monolayer featured in Figure 2) is covalently bound to the sensor surface (i.e., Au and Sulfur of the monolayer), and wherein the sensor is capable of electrochemically detecting a specific binding between the recognition molecules and an analyte as can be seen in the electrochemical teachings on page 5993 ( $1^{st}$  ¶ and Rt column  $2^{nd}$  ¶).

With regard to claim 22, Yousaf et al. teach that a cyclic voltammetry was performed with a Bio-analytical Systems (CV-50 potentiostat by using the gold/SAM as the working electrode, platinum wire as the counter electrode, and Ag/AgCl as the reference electrode (Materials and Methods, Electrochemsitry, Pg.5993).

localized).

substrate.

With regard to claims 23 and 24, Yousaf et al. teach that each of the plurality of patterned, localized micro-electronically addressable sensors are individually activatable and addressable (i.e., via application of electrical potential and subsequent Diels Alder mediated

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With regard to claims 25 and 26, Yousaf et al. teach that the sensor surface includes a chemical molecule based, SAM anchoring layer (Figure 2) covalently bonded to the gold

immobilization of peptide on right of Figure 2 and Figure 1 for teaching of patterned and

With regard to claim 27, Yousaf et al. teach that the SAM anchoring layer is activatable via an application of an electrical potential of 500mV (figure 2).

With regard to claims 28 and 29, Yousaf et al. teach that the electric potential based activation element converts the once inert monolayer (SAM) to a monolayer presenting the corresponding quinine groups which ultimately results in their Diels-Alder mediated immobilization of peptides (Furthest right in Figure 2).

With regard to claim 30, Yousaf et al. teach that the material is gold which is a transition metal.

With regard to claims 31 and 32, Yousaf et al. teach that the thermal activation element is a potentiostat (CV-50) which includes a resistor.

With regard to claim 41, Yousaf et al. teach that their device is a microelectronic chip considering it operates using micro-scaled features that rely on electronic conductance (Fig.1).

With regard to claim 50, Yousaf et al. teach a method for electrochemically detecting (Page 5993 1<sup>st</sup> ¶ and Rt column 2<sup>nd</sup> ¶), a binding event, the method comprising: providing a

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sensing device according to claim 21 as can be seen above, activating the sensor surface; depositing a recognition molecule from a liquid phase or a vapor phase onto the sensor surface (see above rejection for claims 21-32); and detecting a binding event between the recognition molecule and an analyte (i.e., detection was measured by scanning electron micrograph and via fluorescence microscopy (Figures 3-5 page 5995).

With regard to claims 21 and 50-59 in light of the above indefiniteness issue, Yousaf et al. does not teach that the part of the anchoring layer that is either heated or cooled or the exact size of this area.

Hollis et al. teaches the use of embedded resistors (32) described in connection with FIGS. 1 and 4 to locally heat predetermined array test sites without substantially heating adjacent sites (Col. 13 lines 50-60). Furthermore, Hollis et al. teaches that about 7+million test sites can be fabricated and tested on a single 3 in. Silicon wafer using the state of technology at the time when this patent was issued.

At the time the invention was made it would have been obvious to a person of skill in the art to have incorporated the regions and heaters sized as taught by Hollis et al. into the device of Yousaf et al. as the ability to affect multiple changes in the bioactivity of a substrate will be especially useful in studies of heterotypic cell-cell interactions where the fate of a given cell depends on the identities and periods of exposure to neighboring cells. Such dynamic substrate would also prove useful in microfluidic lab-on-a-chip systems by allowing active use of channels to process analytes in a sample. In addition Hollis provides that the heating of each selected region results in the ability to deprotect (revealing free -OH groups) at selected locations making the addition of nucleic acid bases a more specific venture (Col. 12).

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2. Claims 60-62 are rejected under 35 U.S.C. 103(a) as being unpatentable over Yousaf et

al.(PNAS, 2001) in view Hollis et al. (US 5653939) and in further view of Shin et al. (Applied

Surface Science 214(2003) 214-221).

The teachings of Yousaf in view of Hollis et al. can be seen above.

With respect to claim 63, Yousaf et al. teach the use of hydroquinone-terminated

alkanethiol (HQ) and penta (ethylene glycol) terminated alkanethiol (EG5OH) to install the

electroactive monolayer in the nonprinted regions.

Yousaf in view of Hollis et al. do not teach an ISFET or the use of Ta205.

Shin et al. teach tantalum pentoxide for use as the pH sensitive layer for ISFET

(Introduction page 214).

It would have been obvious to a person of skill in the art at the time the invention was

made to have include the ISFET and Ta2O5 of Shin into the device of Yousaf in view of Hollis

as Shin teaches that Ta2O5 shows the best pH sensitive properties in terms of sensitively and

stability making it a superior choice for cost, efficiency and resource savings. Furthermore, it is

well known that it is obvious to combine prior art elements according to known methods to yield

predictable results.

**Response to Arguments** 

Applicant's arguments with respect to claims 21-32 have been considered but are moot in

view of the new ground(s) of rejection.

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In the interest of compact prosecution, the applicant is encourage to include positively recited structural limitations of their sensor, its structural constitution comprised by various elements and recitations of the proximity and position of each of these elements to one another in order to obviate the presently cited art. The applicant is encouraged to phone the examiner with their ideas should they desire feedback before their submission.

#### Conclusion

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Sally A. Sakelaris whose telephone number is 5712726297. The examiner can normally be reached on Monday-Friday 8-5:00.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Jill Warden can be reached on 5712721267. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Sally A Sakelaris/

Examiner, Art Unit 1773

4/26/2011